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ANSWER 1 OF 1 REGISTRY COPYRIGHT 2008 ACS on STN
L5
RN
     703404-85-7 REGISTRY
ED
     Entered STN: 04 Jul 2004
     9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-
CN
     i][1,6]benzodiazocine-10-carboxylic acid,
     10-amino-2, 3, 9, 10, 11, 12-hexahydro-1, 3-dioxo-, 2-aminoethyl ester,
     (9R, 10S, 12S) - rel - (9CI) (CA INDEX NAME)
     STEREOSEARCH
FS
     C28 H23 N5 O4
MF
CI
     COM
SR
     CA
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Relative stereochemistry.

PROPERTY DATA AVAILABLE IN THE 'PROP' FORMAT

L6 ANSWER 1 OF 8 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2008:271655 CAPLUS

DOCUMENT NUMBER: 148:535309

TITLE: Tau kinase inhibitors protect hippocampal synapses

despite of insoluble tau accumulation

AUTHOR(S): Hinners, Ina; Hill, Anika; Otto, Ulrike; Michalsky,

Anke; Mack, Till G. A.; Striggow, Frank

CORPORATE SOURCE: KeyNeurotek Pharmaceuticals AG, Magdeburg, D-39120,

Germany

SOURCE: Molecular and Cellular Neuroscience (2008), 37(3),

559-567

CODEN: MOCNED; ISSN: 1044-7431

PUBLISHER: Elsevier
DOCUMENT TYPE: Journal
LANGUAGE: English

AB A better understanding of the cellular and mol. pathomechanisms of Alzheimer's disease (AD) is a prerequisite for the development of efficient treatments. We have used a novel assay system based on virus-transduced organotypic hippocampal slice cultures that mimics important aspects of tau-driven AD pathol. in a short time frame. Human tau P301L, when expressed in pyramidal neurons of hippocampal slice cultures, was increasingly phosphorylated at several disease-relevant epitopes, leading to progressive neuronal dystrophy and formation of RIPA-insol. tau. AD-like tau hyperphosphorylation was reduced by the tau kinase inhibitors lithium and SRN-003-556, but RIPA-insol. tau remained unaffected after treatment with any of these substances. Only SRN-003-556 was able to protect hippocampal neurons from synaptic damage that was presumably caused by a toxic soluble tau fraction. These data provide first mechanistic insights towards the functional benefits of SRN-003-556 that have been observed in vivo.

IT 906665-84-7, SRN 003-556

RL: BUU (Biological use, unclassified); PAC (Pharmacological activity); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(tau kinase inhibitors protect hippocampal synapses despite of insol. tau accumulation)

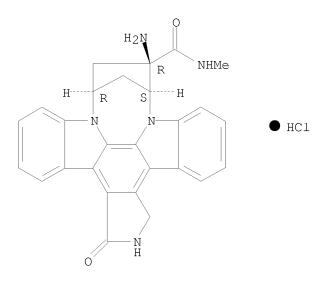
RN 906665-84-7 CAPLUS

CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-

i][1,6]benzodiazocine-10-carboxamide,

10-amino-2,3,9,10,11,12-hexahydro-N-methyl-1-oxo-, hydrochloride (1:1), (9S,10R,12R)- (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 38 THERE ARE 38 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ACCESSION NUMBER:

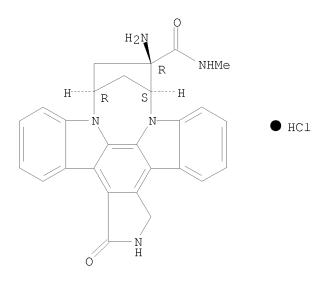
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DOCUMENT NUMBER:
                         146:514513
TITLE:
                         Development of inhibitors of pathological tau
                         hyperphosphorylation for a disease modifying approach
                         to Alzheimer's disease
AUTHOR(S):
                         Sahagun, H.; Huebinger, G.; Klafki, H.; Gordon, S.;
                         Ferrand, S.; Muehlbacher, S.; Mueller, S.; Seneci, P.;
                         Monse, B.; Froehner, W.; Casiraghi, L.; du Boullay, O.
                         Thillaye; Braxmeier, T.; LeCorre, S.; Dirscherl, L.;
                         Jolivalt, C.; Lahu, G.; Sobek-Klocke, I.; Plesnila,
                         N.; Hutton, M.; Roder, H.
CORPORATE SOURCE:
                         Sirenade Pharmaceuticals AG, Munich, Germany
SOURCE:
                         New Trends in Alzheimer and Parkinson Related
                         Disorders: ADPD 2005, Collection of Selected Free
                         Papers of the International Conference on Progress in
                         Alzheimer's and Parkinson's Disease (AD/PD), 7th,
                         Sorrento, Italy, Mar. 9-13, 2005 (2005), 71-76.
                         Editor(s): Fisher, A. Monduzzi Editore: Bologna,
                         Italy.
                         CODEN: 69IYJM; ISBN: 88-7587-174-4
DOCUMENT TYPE:
                         Conference
LANGUAGE:
                         English
     Pathol. hyperphosphorylation of tau protein appears to be an early and
     critical step in the process leading to tau-pathol. and neurodegeneration in
     Alzheimer's disease (AD) and other tauopathies. Pathol.
     hyperphosphorylation of tau is believed to be the consequence of an
     imbalance of kinases and phosphatases that results in the phosphorylation
     of critical sites, leading to detachment from microtubules, accumulation in
     the somatodendritic compartment and ultimately aggregation to form paired
     helical filaments (PHFs). One possible means to modify the course of AD
     and other tauopathies is to inhibit the putative kinase activities leading
     to the inappropriate phosphorylation of tau. Based on initial in vitro
     evidence related to the maximal stoichiometry of tau phosphorylation, the
     mitogen activated/extracellular signal regulated kinase MAPK/ERK2 was
     considered for developing a targeted library of compds. with kinase
     inhibitory activities. Promising inhibitor candidates were identified in
     a screening cascade comprised of primary kinase assays, cellular models of
     PHF-type tau hyperphosphorylation, and a transgenic animal model of
     authentic neurofibrillary degeneration.
ΙT
     906665-84-7
     RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (SRN 003-556; SRN-003-556 inhibited human tau hyperphosphorylation in
        hippocampal rat model of Alzheimer's disease)
     906665-84-7 CAPLUS
RN
     9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-
CN
     i][1,6]benzodiazocine-10-carboxamide,
     10-amino-2,3,9,10,11,12-hexahydro-N-methyl-1-oxo-, hydrochloride (1:1),
     (9S, 10R, 12R) - (CA INDEX NAME)
```

ANSWER 2 OF 8 CAPLUS COPYRIGHT 2008 ACS on STN

2007:232200 CAPLUS

Absolute stereochemistry.



REFERENCE COUNT: 13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

ANSWER 3 OF 8 CAPLUS COPYRIGHT 2008 ACS on STN 1.6

ACCESSION NUMBER: 2006:647437 CAPLUS

DOCUMENT NUMBER: 145:263072

TITLE: An inhibitor of tau hyperphosphorylation prevents

severe motor impairments in tau transgenic mice

Le Corre, Sylvie; Klafki, Hans W.; Plesnila, Nikolaus; AUTHOR(S):

Huebinger, Gabriele; Obermeier, Axel; Sahagun, Heidi; Monse, Barbara; Seneci, Pierfausto; Lewis, Jada; Eriksen, Jason; Zehr, Cynthia; Yue, Mei; McGowan, Eileen; Dickson, Dennis W.; Hutton, Michael; Roder,

Hanno M.

CORPORATE SOURCE:

Sirenade Pharmaceuticals, Martinsried, 82152, Germany Proceedings of the National Academy of Sciences of the SOURCE:

United States of America (2006), 103(25), 9673-9678

CODEN: PNASA6; ISSN: 0027-8424

PUBLISHER: National Academy of Sciences

DOCUMENT TYPE: Journal LANGUAGE: English

An orally bioavailable and blood-brain barrier penetrating analog of the kinase inhibitor K252a was able to prevent the typical motor deficits in the tau (P301L) transgenic mouse model (JNPL3) and markedly reduce soluble aggregated hyperphosphorylated tau. However, neurofibrillary tangle counts were not reduced in the successfully treated cohort, suggesting that the main cytotoxic effects of tau are not exerted by neurofibrillary tangles but by lower mol. mass aggregates of tau. Our findings strongly suggest that abnormal tau hyperphosphorylation plays a critical role in the development of tauopathy and suggest a previously undescribed treatment strategy for neurodegenerative diseases involving tau pathol.

ΙT 906665-84-7, SRN 003-556

> RL: PAC (Pharmacological activity); PKT (Pharmacokinetics); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(inhibitor of tau hyperphosphorylation prevents severe motor impairments in tau transgenic mice)

906665-84-7 CAPLUS RN

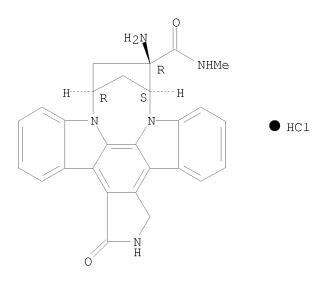
9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-CN

i][1,6]benzodiazocine-10-carboxamide,

10-amino-2,3,9,10,11,12-hexahydro-N-methyl-1-oxo-, hydrochloride (1:1),

(9S, 10R, 12R) - (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 26 THERE ARE 26 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

PUBLISHER:

L6 ANSWER 4 OF 8 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2005:699415 CAPLUS

DOCUMENT NUMBER: 143:347145

TITLE: The synthesis of bioactive indolocarbazoles related to

K-252a

AUTHOR(S): Moffat, David; Nichols, Christopher J.; Riley, Dean

A.; Simpkins, Nigel S.

CORPORATE SOURCE: Celltech Therapeutics Ltd, Slough, SL1 4EN, UK SOURCE: Organic & Biomolecular Chemistry (2005), 3(16),

2953-2975

CODEN: OBCRAK; ISSN: 1477-0520 Royal Society of Chemistry

DOCUMENT TYPE: Journal LANGUAGE: English

OTHER SOURCE(S): CASREACT 143:347145

AB A range of functionalized indolocarbazoles, related to the natural product K-252a, have been prepared, starting from a readily available bridged cyclopentene, cis-3,5-dibromocyclopentene. Sequences of transformations, involving initial hydroboration-oxidation to give a ketone, or by dihydroxylation and cyclic sulfate formation, enable the preparation of diverse indolocarbazole products. Issues of imide nitrogen protection for the indolocarbazole, and opportunities for asym. desymmetrization of key intermediates were also explored. A novel chiral lithium amide base mediated transformation of a cyclic sulfate intermediate gave the anticipated ketone product in up to 87% ee. A number of compds., in the form of unprotected imide substituted indolocarbazoles, were screened for biol. activity and were found to be potent inhibitors of a number of kinase enzymes.

IT 703404-92-6P

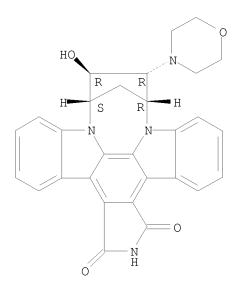
RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent)

(preparation of functionalized indolocarbazole derivs. and study of its activity as protein kinase inhibitors)

RN 703404-92-6 CAPLUS

CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
10-azido-9,10,11,12-tetrahydro-11-hydroxy-, (9R,10R,11R,12S)-rel- (9CI)
(CA INDEX NAME)

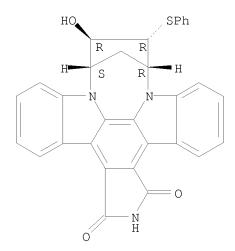
Relative stereochemistry.



RN 865485-75-2 CAPLUS CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3(2H)-dione, 9,10,11,12-tetrahydro-10-hydroxy-11-(phenylthio)-, (9R,10S,11S,12S)-rel-

(9CI) (CA INDEX NAME)

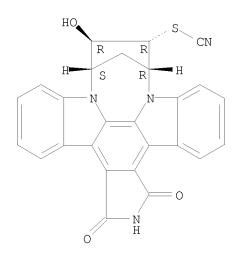
Relative stereochemistry.



RN 865485-76-3 CAPLUS

CN Thiocyanic acid, (9R,10R,11R,12S)-2,3,9,10,11,12-hexahydro-11-hydroxy-1,3-dioxo-9,12-methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocin-10-yl ester, rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.



RN 865485-77-4 CAPLUS

CN 1H-1,2,3-Triazole-4,5-dicarboxylic acid, 1-[(9R,10R,11R,12S)-2,3,9,10,11,12-hexahydro-11-hydroxy-1,3-dioxo-9,12-methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocin-10-yl]-, dimethyl ester, rel- (9CI) (CA INDEX NAME)

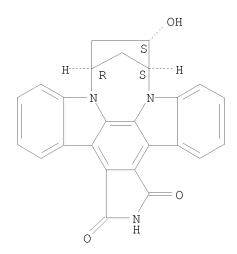
IT 233253-34-4P 233253-35-5P

RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation); RACT (Reactant or reagent)

(preparation of functionalized indolocarbazole derivs. and study of their activity as protein kinase inhibitors)

RN 233253-34-4 CAPLUS

Relative stereochemistry.



RN 233253-35-5 CAPLUS

CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-

i][1,6]benzodiazocine-1,3,10(2H,9H)-trione, 11,12-dihydro- (9CI) (CA INDEX NAME)

IT 233253-37-7P 865606-88-8P

RL: PAC (Pharmacological activity); SPN (Synthetic preparation); BIOL (Biological study); PREP (Preparation)

(preparation of functionalized indolocarbazole derivs. and study of their activity as protein kinase inhibitors)

RN 233253-37-7 CAPLUS

CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-

i][1,6]benzodiazocine-10-carboxylic acid,

2,3,9,10,11,12-hexahydro-10-hydroxy-1,3-dioxo-, methyl ester,

(9R, 10S, 12S) -rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

RN 865606-88-8 CAPLUS

CN 6,10-Methano-16H-[1,3,2]dioxathiolo[4,5-c]diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-16,18(17H)-dione, 6,6a,9a,10-tetrahydro-, 8-oxide, (6R,6aS,9aR,10S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

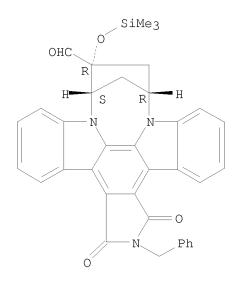
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233253-26-4P 233253-27-5P 233253-30-0P
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     703405-34-9P 786688-05-9P 786688-06-0P
     786688-07-1P 786688-09-3P 786688-10-6P
     786688-12-8P 786688-13-9P 865485-69-4P
     865485-72-9P 865485-73-0P 865606-87-7P
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (preparation of functionalized indolocarbazole derivs. related to natural
       product K-252a using indolo[2,3-a]pyrrolo[3,4-c]carbazole dione and
        cis-di(bromo)cyclopentene as starting materials)
RN
     233253-26-4 CAPLUS
     9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-
CN
     i][1,6]benzodiazocine-1,3(2H)-dione,
     9,10,11,12-tetrahydro-10-hydroxy-2-(phenylmethyl)-, (9R,10R,12S)-rel-
     (9CI) (CA INDEX NAME)
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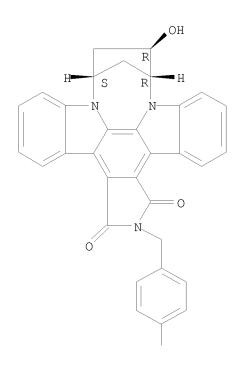
RN 233253-27-5 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3,10(2H,9H)-trione,
11,12-dihydro-2-(phenylmethyl)- (9CI) (CA INDEX NAME)

RN 233253-30-0 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
10-ethenyl-9,10,11,12-tetrahydro-2-(phenylmethyl)-10-[(trimethylsilyl)oxy], (9R,10R,12S)-rel- (9CI) (CA INDEX NAME)

RN 233253-31-1 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carboxaldehyde,
2,3,9,10,11,12-hexahydro-1,3-dioxo-2-(phenylmethyl)-10[(trimethylsilyl)oxy]-, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.



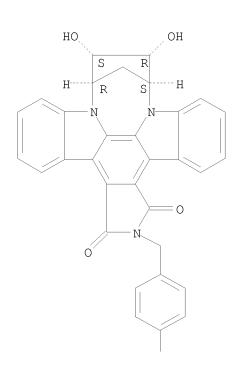


PAGE 2-A | OMe

RN 253680-58-9 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3,10(2H,9H)-trione,
11,12-dihydro-2-[(4-methoxyphenyl)methyl]- (9CI) (CA INDEX NAME)

RN 703405-34-9 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-10,11-dihydroxy-2-[(4-methoxyphenyl)methyl]-,
(9R,10S,11R,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.



PAGE 1-A

PAGE 2-A

RN 786688-05-9 CAPLUS

CN 6,10-Methano-16H-[1,3,2]dioxathiolo[4,5-c]diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-16,18(17H)-dione, 6,6a,9a,10-tetrahydro-17-[(4-methoxyphenyl)methyl]-, 8,8-dioxide, (6R,6aS,9aR,10S)-rel- (9CI) (CA INDEX NAME)

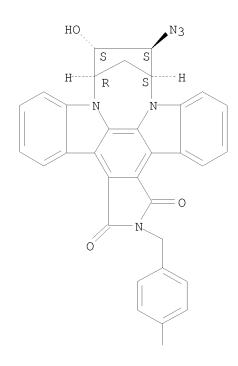
Relative stereochemistry.

PAGE 1-A

PAGE 2-A

RN 786688-06-0 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
 10-azido-9,10,11,12-tetrahydro-11-hydroxy-2-[(4-methoxyphenyl)methyl]-,
 (9R,10R,11R,12S)-rel- (9CI) (CA INDEX NAME)

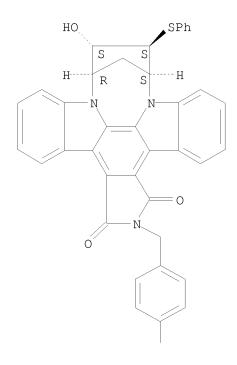
Relative stereochemistry.



PAGE 1-A

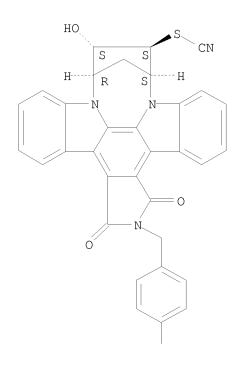
PAGE 2-A | OMe

PAGE 2-A



PAGE 2-A | OMe

RN 786688-10-6 CAPLUS
CN Thiocyanic acid, (9R,10R,11R,12S)-2,3,9,10,11,12-hexahydro-11-hydroxy-2-[(4-methoxyphenyl)methyl]-1,3-dioxo-9,12-methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocin-10-yl ester, rel- (9CI) (CA INDEX NAME)



PAGE 2-A | OMe

RN 786688-12-8 CAPLUS
CN 1H-1,2,3-Triazole-4,5-dicarboxylic acid,
1-[(9R,10R,11R,12S)-2,3,9,10,11,12-hexahydro-11-hydroxy-2-[(4-methoxyphenyl)methyl]-1,3-dioxo-9,12-methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocin-10-yl]-, dimethyl ester, rel- (9CI)
(CA INDEX NAME)

PAGE 2-A

RN 786688-13-9 CAPLUS

CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3,10(2H,9H)-trione,
11,12-dihydro-2-[(4-methoxyphenyl)methyl]-, (9S,12R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

PAGE 2-A | OMe

RN 865485-69-4 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
10-ethenyl-9,10,11,12-tetrahydro-10-hydroxy-2-(phenylmethyl)-,
(9R,10R,12S)-rel- (9CI) (CA INDEX NAME)

RN 865485-72-9 CAPLUS

CN Acetic acid, trifluoro-, (9R,10S,11S,12S)-11-azido-2,3,9,10,11,12-hexahydro-1,3-dioxo-9,12-methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocin-10-yl ester, rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

RN 865485-73-0 CAPLUS

CN Acetic acid, trifluoro-, (9R,10S,11S,12S)-2,3,9,10,11,12-hexahydro-1,3-dioxo-11-(phenylthio)-9,12-methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocin-10-yl ester, rel- (9CI) (CA INDEX NAME)

RN

865606-87-7 CAPLUS 6,10-Methano-16H-[1,3,2]dioxathiolo[4,5-c]diindolo[1,2,3-fg:3',2',1'-k1]pyrrolo[3,4-i][1,6]benzodiazocine-16,18(17H)-dione, CN 6,6a,9a,10-tetrahydro-17-[(4-methoxyphenyl)methyl]-, 8-oxide, (6R,6aS,9aR,10S)-rel- (9CI) (CA INDEX NAME)

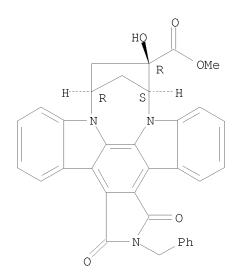
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PAGE 2-A

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     786688-14-0P 865485-68-3P 865485-70-7P
     865485-71-8P 865485-80-9P 865485-81-0P
     865485-82-1P 865485-83-2P 865485-84-3P
     RL: SPN (Synthetic preparation); PREP (Preparation)
        (preparation of functionalized indolocarbazole derivs. related to natural
       product K-252a using indolo[2,3-a]pyrrolo[3,4-c]carbazole dione and
        cis-di(bromo)cyclopentene as starting materials)
     233253-28-6 CAPLUS
RN
     9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-
CN
     i][1,6]benzodiazocine-10-carbonitrile,
     2,3,9,10,11,12-hexahydro-1,3-dioxo-2-(phenylmethyl)-10-
     [(trimethylsilyl)oxy]-, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)
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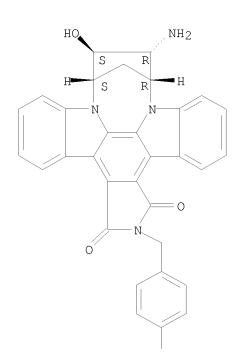
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CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-10-carboxylic acid,
2,3,9,10,11,12-hexahydro-10-hydroxy-1,3-dioxo-2-(phenylmethyl)-, methyl ester, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.



RN 703404-93-7 CAPLUS CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3(2H)-dione, 10-amino-9,10,11,12-tetrahydro-11-hydroxy-, (9R,10R,11S,12S)-rel- (9CI) (CA INDEX NAME)

RN 703405-10-1 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
10-amino-9,10,11,12-tetrahydro-11-hydroxy-2-[(4-methoxyphenyl)methyl]-,
(9R,10R,11S,12S)-rel- (9CI) (CA INDEX NAME)



PAGE 2-A | OMe

RN 786688-08-2 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-k1]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
 10-(benzoyloxy)-9,10,11,12-tetrahydro-11-hydroxy-2-[(4methoxyphenyl)methyl]-, (9R,10R,11R,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

PAGE 1-A

PAGE 2-A | OMe

PAGE 2-A



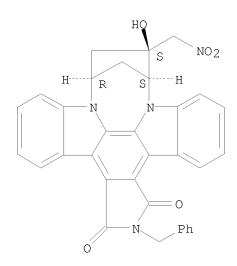
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PAGE 2-A | OMe

RN 865485-68-3 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-10-hydroxy-2-(phenylmethyl)-, (9R,10S,12S)-rel(9CI) (CA INDEX NAME)

RN 865485-70-7 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-k1]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-10-hydroxy-10-(nitromethyl)-2-(phenylmethyl)-,
(9R,10R,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.



RN 865485-71-8 CAPLUS
CN 6,8-Methano-14H-diindolo[1,2,3-fg:3',2',1'-kl]oxireno[c]pyrrolo[3,4-i][1,6]benzodiazocine-14,16(15H)-dione,
6,6a,7a,8-tetrahydro-15-(phenylmethyl)-, (6R,6aS,7aR,8S)-rel- (9CI) (CA INDEX NAME)

RN 865485-80-9 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3,10(2H,9H)-trione,
11,12-dihydro-2-[(4-methoxyphenyl)methyl]-11,11-di-2-propenyl- (9CI) (CAINDEX NAME)

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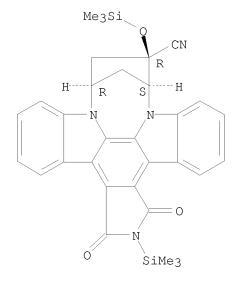
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CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3,10(2H,9H)-trione,
11,12-dihydro-2-[(4-methoxyphenyl)methyl]-11,11-dimethyl- (9CI) (CA INDEX NAME)

RN 865485-82-1 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3,10(2H,9H)-trione,
11,12-dihydro-2-[(4-methoxyphenyl)methyl]-11,11-bis(phenylmethyl)- (9CI)
(CA INDEX NAME)

RN 865485-83-2 CAPLUS
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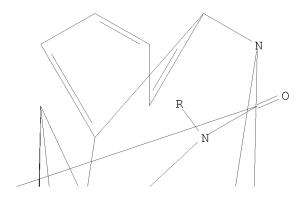
2,3,9,10,11,12-hexahydro-1,3-dioxo-2-(trimethylsilyl)-10-[(trimethylsilyl)oxy]-, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.



RN 865485-84-3 CAPLUS

CN 6,8-Methano-14H-diindolo[1,2,3-fg:3',2',1'-kl]oxireno[c]pyrrolo[3,4-i][1,6]benzodiazocine-14,16(15H)-dione,
6,6a,7a,8-tetrahydro-15-[(4-methoxyphenyl)methyl]-, (6R,6aS,7aR,8S)-rel-(9CI) (CA INDEX NAME)



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- * STRUCTURE DIAGRAM TOO LARGE FOR DISPLAY AVAILABLE VIA OFFLINE PRINT *
 REFERENCE COUNT: 85 THERE ARE 85 CITED REFERENCES AVAILABLE FOR THIS
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10/532,263

L6 ANSWER 5 OF 8 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2004:745005 CAPLUS

DOCUMENT NUMBER: 141:395536

TITLE: Synthesis of bioactive indolocarbazoles: synthesis,

nucleophilic ring-opening and chiral base

desymmetrisation of a cyclic sulfate intermediate

AUTHOR(S): Nichols, Christopher J.; Simpkins, Nigel S. CORPORATE SOURCE: School of Chemistry, University of Nottingham,

University Park, NG7 2RD, UK

SOURCE: Tetrahedron Letters (2004), 45(40), 7469-7473

CODEN: TELEAY; ISSN: 0040-4039

PUBLISHER: Elsevier B.V.

DOCUMENT TYPE: Journal LANGUAGE: English

OTHER SOURCE(S): CASREACT 141:395536

GΙ

AB A number of new functionalized bridged indolocarbazoles I (R1 = 4-MeOC6H4CH2; R2 = N3, H2N, 4-morpholinyl, PhCO2, PhS, SCN, etc.) has been prepared by ring-opening reactions of a key cyclic sulfate intermediate I (R2R3 = OSO2O), prepared from the corresponding diol (R2 = R3 = HO) by treatment with sulfuryl diimidazole and DBU. The same cyclic sulfate also undergoes an unprecedented asym. rearrangement to a chiral ketone II, on treatment with a chiral lithium amide base.

IT 703405-34-9P 786688-05-9P 786688-06-0P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(preparation of functionalized bridged indolocarbazoles via nucleophilic ring-opening and chiral base desymmetrisation of cyclic sulfate derivative) 703405-34-9 CAPLUS

RN 703405-34-9 CAPLUS CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-

i][1,6]benzodiazocine-1,3(2H)-dione,

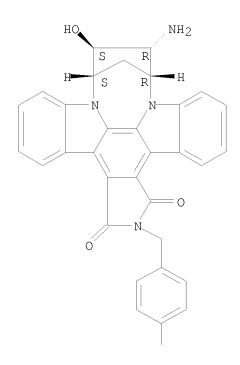
9, 10, 11, 12-tetrahydro-10, 11-dihydroxy-2-[(4-methoxyphenyl)methyl]-,

(9R,10S,11R,12S)-rel- (9CI) (CA INDEX NAME)

PAGE 2-A | OMe

RN 786688-05-9 CAPLUS
CN 6,10-Methano-16H-[1,3,2]dioxathiolo[4,5-c]diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-16,18(17H)-dione,
6,6a,9a,10-tetrahydro-17-[(4-methoxyphenyl)methyl]-, 8,8-dioxide,
(6R,6aS,9aR,10S)-rel- (9CI) (CA INDEX NAME)

PAGE 2-A | OMe



PAGE 2-A | OMe

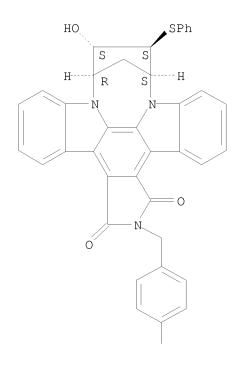
RN 786688-07-1 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
 9,10,11,12-tetrahydro-10-hydroxy-2-[(4-methoxyphenyl)methyl]-11-(4 morpholinyl)-, (9R,10S,11S,12S)-rel- (9CI) (CA INDEX NAME)

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RN 786688-08-2 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
 10-(benzoyloxy)-9,10,11,12-tetrahydro-11-hydroxy-2-[(4 methoxyphenyl)methyl]-, (9R,10R,11R,12S)-rel- (9CI) (CA INDEX NAME)

PAGE 2-A | OMe

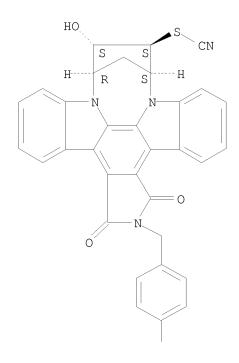
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CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
 9,10,11,12-tetrahydro-10-hydroxy-2-[(4-methoxyphenyl)methyl]-11 (phenylthio)-, (9R,10S,11S,12S)-rel- (9CI) (CA INDEX NAME)



PAGE 2-A | OMe

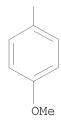
RN 786688-10-6 CAPLUS

CN Thiocyanic acid, (9R,10R,11R,12S)-2,3,9,10,11,12-hexahydro-11-hydroxy-2-[(4-methoxyphenyl)methyl]-1,3-dioxo-9,12-methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocin-10-yl ester, rel- (9CI) (CA INDEX NAME)



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RN 786688-12-8 CAPLUS
CN 1H-1,2,3-Triazole-4,5-dicarboxylic acid,
 1-[(9R,10R,11R,12S)-2,3,9,10,11,12-hexahydro-11-hydroxy-2-[(4-methoxyphenyl)methyl]-1,3-dioxo-9,12-methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocin-10-yl]-, dimethyl ester, rel- (9CI)
 (CA INDEX NAME)

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RN 786688-13-9 CAPLUS

CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3,10(2H,9H)-trione, 11,12-dihydro-2-[(4-methoxyphenyl)methyl]-, (9S,12R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry. Rotation (-).

PAGE 2-A | OMe

Absolute stereochemistry. Rotation (-).

PAGE 2-A

OMe

REFERENCE COUNT: 24 THERE ARE 24 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 6 OF 8 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2004:467897 CAPLUS

DOCUMENT NUMBER: 141:38635

TITLE: Preparation of N,N-bridged, nitrogen-substituted

carbacyclic indolocarbazoles for use in pharmaceutical

compositions as protein kinase inhibitors

INVENTOR(S): Monse, Barbara; Braxmeier, Tobias; Ferrand, Sandrine;

Gordon, Sandra; Klafki, Hans; Lahu, Gezim; Roder, Hanno; Sahagun-Krause, Heidi; Seneci, Pierfausto;

APPLICATION NO

DATE

Thillaye du Boullay, Olivier

PATENT ASSIGNEE(S): Nad A.-G., Germany SOURCE: PCT Int. Appl., 89 pp.

CODEN: PIXXD2

KIND DATE

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA.	PAIENI NO.					KIND DAIE			APPLICATION NO.						DAIE			
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		TR,	BF,	ВJ,	CF,	CG,	CI,	CM,	GΑ,	GN,	GQ,	GW,	ML,	MR,	NE,	SN,	TD,	ΤG
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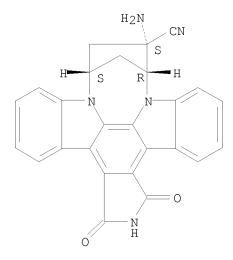
GΙ

AB This invention relates to the preparation of novel carbacyclic indolocarbazoles, such as I [R1 = NR13R14; R2 = H, CN, alkyl, aryl, heteroaryl, acyl, carboxy, carboxamido; R1R2 = spiro nitrogen containing heterocycle, such as spirohydantoyl; R3 = H, OR13, OCOR13, OCONHR13, OCONR13R14; R1R3 = fused heterocycle, such as -OSO20-, and R2 = H; R4, R6 = H, CN, halogen, alkyl, alkenyl, alkynyl, aryl, heteroaryl, alkoxy, acyl, carboxy. carboxamido, etc.; R8R9, R10R11 = H2, O, S; R8 = H, R9 = OH; R10 = H, R11 = OH; R12 = H, alkyl, cycloalkyl, benzyl aryl heteroaryl, acyl, carboxy, etc.; R13, R14 = H, alkyl, cycloalkyl, acyl, aryl, etc.], for therapeutic use as protein kinase inhibitors with advantageous pharmaceutical properties. These indolocarbazoles are claimed for use in the treatment of CNS diseases, non-insulin-dependent diabetes mellitus, acute stroke and other neuro-traumatic injuries, diabetes mellitus, malignant diseases, diseases caused by malfunctioning of specific signaling pathways and neurodegenerative diseases, such as Alzheimer's disease. Thus, indolocarbazole II (R1 = β -NH2, R2 = α -H, R3 = H) was prepared starting from cyclopentadiene, 4-methoxybenzyl amine, dichloromaleic anhydride, and indole via a multistep synthetic sequence which included a reaction of 12,13-dihydro-6-[(4-methoxyphenyl)methyl]-5Hindolo[2,3-a]pyrrolo[3,4-c]carbazole-5,7(6H)-dione with cis-3,5-dibromocyclopentene using NaH in THF to form II (R1R3 = bond, R2 = H) in 90.4% yield. II (R1R3 = bond, R2 = H), which contains the target ring skeleton, further underwent an hydroxylation sequence using BH2.THF followed by NaOH and H2O2 to form alc. II (R1 = α -OH, R2 = β -H, R3 = H), oxidation of the alc. to the corresponding ketone II (R1R2 = 0, R3 = H), reaction of the ketone with benzylamine to give N-benzyl amine II (R1 = β -NHCH2Ph, R2 = α -H, R3 = H) and, finally, N-debenzylation to give the desired indolocarbazole II (R1 = β -NH2, R2 = α -H, R3 = H). The prepared indolocarbazoles were assayed for inhibiting the activity of a group of protein kinases consisting of extracellular signal regulated kinase 2 (ERK2), protein kinase A (PKA), protein kinase C (PKC) and glycogen synthase kinase 3β (GSK3 β). ΙT 703404-81-3P 703404-82-4P 703404-84-6P 703404-90-4P 703404-92-6P 703404-94-8P 703404-97-1P 703404-98-2P 703404-99-3P

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     704915-60-6P, NAD 0241
     RL: PAC (Pharmacological activity); RCT (Reactant); SPN (Synthetic
     preparation); THU (Therapeutic use); BIOL (Biological study); PREP
     (Preparation); RACT (Reactant or reagent); USES (Uses)
        (preparation of N, N-bridged, nitrogen-substituted carbacyclic
        indolocarbazoles for use in pharmaceutical compns. as protein kinase
        inhibitors)
RN
     703404-81-3 CAPLUS
CN
     9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-
     i][1,6]benzodiazocine-10-carbonitrile,
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     INDEX NAME)
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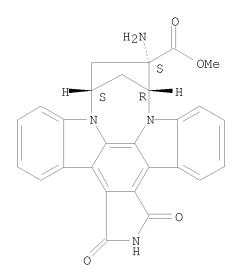
Relative stereochemistry.



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RN 703404-82-4 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-10-carboxylic acid,
10-amino-2,3,9,10,11,12-hexahydro-1,3-dioxo-, ethyl ester,
(9R,10S,12S)-rel- (9CI) (CA INDEX NAME)
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RN 703404-84-6 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-10-carboxylic acid,
10-amino-2,3,9,10,11,12-hexahydro-1,3-dioxo-, methyl ester,
(9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

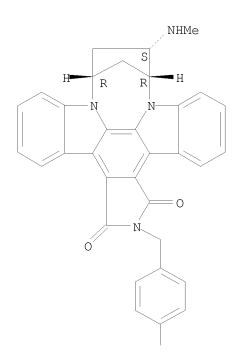


PAGE 2-A

Relative stereochemistry.

RN 703404-97-1 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carboxamide,
10-amino-2,3,9,10,11,12-hexahydro-3-oxo-, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

RN 703404-98-2 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-2-[(4-methoxyphenyl)methyl]-10-(methylamino)-,
(9R,10S,12R)-rel- (9CI) (CA INDEX NAME)



PAGE 2-A | OMe

RN 703404-99-3 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-2-[(4-methoxyphenyl)methyl]-10-[(phenylmethyl)amino], (9R,10S,12R)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

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PAGE 2-A | OMe

RN 703405-03-2 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-2-[(4-methoxyphenyl)methyl]-10-(4-morpholinyl)-,
(9R,10S,12R)-rel- (9CI) (CA INDEX NAME)

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RN 703405-05-4 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carbonitrile,
10-amino-2,3,9,10,11,12-hexahydro-2-[(4-methoxyphenyl)methyl]-1,3-dioxo-,
(9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

PAGE 2-A

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RN 703405-11-2 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-
i][1,6]benzodiazocine-10-carboxamide,
10-amino-2,3,9,10,11,12-hexahydro-2-[(4-methoxyphenyl)methyl]-1,3-dioxo-,
(9R,10S,12S)-rel- (9CI) (CA INDEX NAME)
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PAGE 2-A

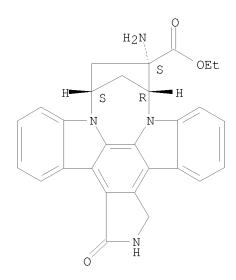
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CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carboxylic acid,
10-amino-2,3,9,10,11,12-hexahydro-2-[(4-methoxyphenyl)methyl]-1,3-dioxo-,
ethyl ester, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

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RN 703405-17-8 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carboxamide,
 10-amino-2,3,9,10,11,12-hexahydro-1-oxo-, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

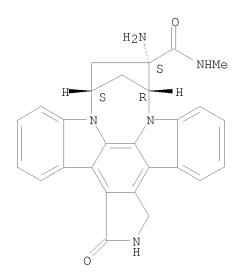
RN 703405-18-9 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carboxylic acid,
10-amino-2,3,9,10,11,12-hexahydro-1-oxo-, ethyl ester, (9R,10S,12S)-rel(9CI) (CA INDEX NAME)

Relative stereochemistry.

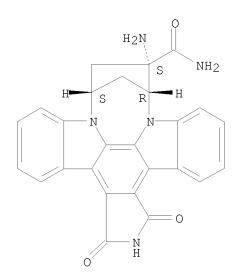


RN 703405-19-0 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carboxylic acid,
10-amino-2,3,9,10,11,12-hexahydro-3-oxo-, ethyl ester, (9R,10S,12S)-rel(9CI) (CA INDEX NAME)

Relative stereochemistry.



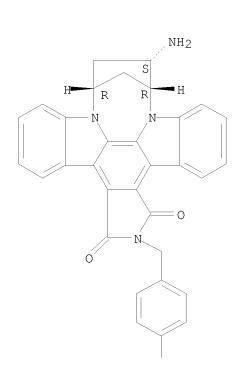
RN 704915-60-6 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carboxamide,
10-amino-2,3,9,10,11,12-hexahydro-1,3-dioxo-, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)



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TT 703404-78-8P 703404-79-9P 703404-80-2P 703404-83-5P 703404-86-8P 703404-87-9P 703404-88-0P 703404-89-1P 703404-91-5P 703404-93-7P 703404-95-9P 703404-96-0P 703405-00-9P 703405-01-0P 703405-02-1P 703405-04-3P 703405-06-5P 703405-10-1P 703405-13-4P 703405-14-5P 703405-15-6P 703405-16-7P 703405-22-5P 703405-23-6P
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10/532,263

Relative stereochemistry.



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RN 703404-79-9 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-10-(methylamino)-, (9R,10S,12R)-rel- (9CI) (CA INDEX NAME)

RN 703404-80-2 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-k1]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-10-(4-morpholinyl)-, (9R,10S,12R)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

RN 703404-83-5 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carboxylic acid,
10-amino-2,3,9,10,11,12-hexahydro-1,3-dioxo-, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

RN 703404-86-8 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-10-carboxylic acid,
10-amino-2,3,9,10,11,12-hexahydro-1,3-dioxo-, 2-aminoethyl ester,
(9R,10S,12S)-rel-, bis(trifluoroacetate) (9CI) (CA INDEX NAME)

CM 1

CRN 703404-85-7 CMF C28 H23 N5 O4

Relative stereochemistry.

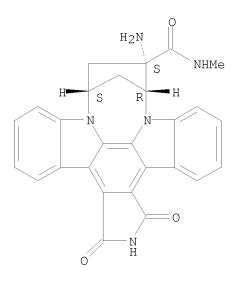
CM 2

CRN 76-05-1

CMF C2 H F3 O2

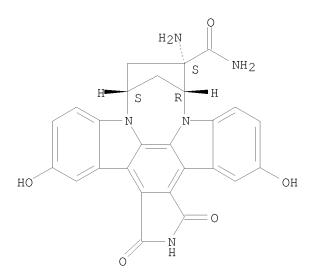
RN 703404-87-9 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carboxamide,
10-amino-2,3,9,10,11,12-hexahydro-N-methyl-1,3-dioxo-, (9R,10S,12S)-rel(9CI) (CA INDEX NAME)

Relative stereochemistry.



RN 703404-88-0 CAPLUS CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-10-carboxamide, 10-amino-2,3,9,10,11,12-hexahydro-N-(2-hydroxyethyl)-1,3-dioxo-, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

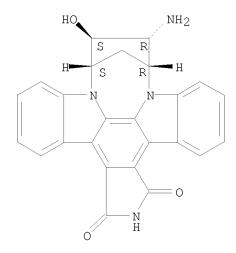
Relative stereochemistry.



RN 703404-91-5 CAPLUS
CN Spiro[imidazolidine-4,10'(9'H)-[9,12]methano[1H]diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine]-1',2,3',5(2'H)-tetrone,
11',12'-dihydro-2'-[(4-methoxyphenyl)methyl]-1-methyl-, (4R,9'S,12'R)-rel-(9CI) (CA INDEX NAME)

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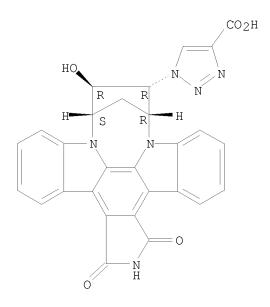
RN 703404-93-7 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
 10-amino-9,10,11,12-tetrahydro-11-hydroxy-, (9R,10R,11S,12S)-rel- (9CI)
 (CA INDEX NAME)



703404-95-9 CAPLUS RN CN 1H-1,2,3-Triazole-4-carboxylic acid, 1-[(9R,10R,11R,12S)-2,3,9,10,11,12-hexahydro-11-hydroxy-1,3-dioxo-9,12-methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocin-

10-yl]-, rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

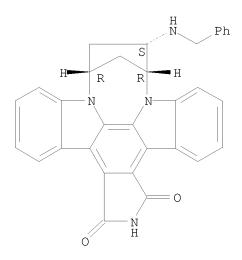


703404-96-0 CAPLUS RN

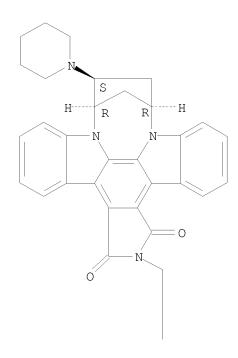
6,10-Methano-16H-diindolo[1,2,3-fg:3',2',1'-kl]oxazolo[4,5-c]pyrrolo[3,4-CN i][1,6]benzodiazocine-8,16,18(9H,17H)-trione, 6,6a,9a,10-tetrahydro-, (6R,6aR,9aR,10S)-rel- (9CI) (CA INDEX NAME)

RN 703405-00-9 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-10-[(phenylmethyl)amino]-, (9R,10S,12R)-rel- (9CI)
(CA INDEX NAME)

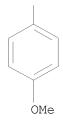
Relative stereochemistry.



RN 703405-01-0 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-2-[(4-methoxyphenyl)methyl]-10-(1-piperidinyl)-,
(9R,10S,12R)-rel- (9CI) (CA INDEX NAME)



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RN 703405-02-1 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-10-[(2-hydroxyethyl)amino]-2-[(4methoxyphenyl)methyl]-, (9R,10S,12R)-rel- (9CI) (CA INDEX NAME)

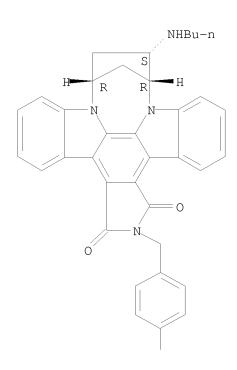
PAGE 2-A | OMe

RN 703405-04-3 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carbonitrile,
2,3,9,10,11,12-hexahydro-2-[(4-methoxyphenyl)methyl]-10-(methylamino)-1,3dioxo-, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

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RN 703405-06-5 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-10-(1-piperidinyl)-, (9R,10S,12R)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.



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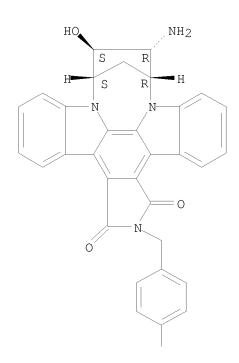
Relative stereochemistry.

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RN 703405-09-8 CAPLUS CN Acetamide, N-[(9R,10S,12S)-10-cyano-2,3,9,10,11,12-hexahydro-1,3-dioxo-9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocin-10-yl]-2,2,2-trifluoro-, rel- (9CI) (CA INDEX NAME)

RN 703405-10-1 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
10-amino-9,10,11,12-tetrahydro-11-hydroxy-2-[(4-methoxyphenyl)methyl]-,
(9R,10R,11S,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.



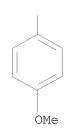
PAGE 1-A

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RN 703405-13-4 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carboxylic acid,
10-amino-2,3,9,10,11,12-hexahydro-2-[(4-methoxyphenyl)methyl]-1,3-dioxo-,
(9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

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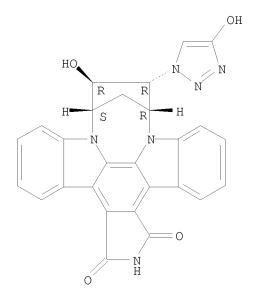
RN 703405-14-5 CAPLUS

CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carboxamide,
10-amino-2,3,9,10,11,12-hexahydro-N,2-bis(2-hydroxyethyl)-1,3-dioxo-,
(9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

RN 703405-15-6 CAPLUS CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3(2H)-dione, 9,10,11,12-tetrahydro-10-hydroxy-11-(4-hydroxy-1H-1,2,3-triazol-1-yl)-, (9R,10S,11S,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.



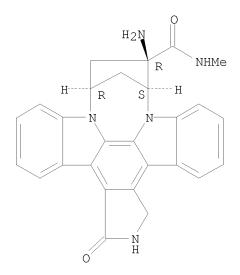
RN 703405-16-7 CAPLUS

CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carboxamide,
10-amino-N-ethyl-2,3,9,10,11,12-hexahydro-1,3-dioxo-, (9R,10S,12S)-rel(9CI) (CA INDEX NAME)

Relative stereochemistry.

RN 703405-22-5 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carboxamide,
10-amino-2,3,9,10,11,12-hexahydro-N-methyl-1-oxo-, (9S,10R,12R)- (9CI)
(CA INDEX NAME)

Absolute stereochemistry.



RN 703405-23-6 CAPLUS CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-

```
i][1,6]benzodiazocine-10-carboxamide,
10-amino-2,3,9,10,11,12-hexahydro-N-methyl-1-oxo-, (9R,10S,12S)- (9CI)
(CA INDEX NAME)
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Absolute stereochemistry.

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RN 703405-25-8 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-10-carboxamide,
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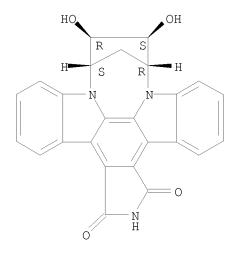
10-amino-2,3,9,10,11,12-hexahydro-N-methyl-3-oxo-, hydrochloride, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

•x HCl

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ΙT
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     253680-58-9P 703405-26-9P 703405-29-2P
     703405-30-5P 703405-31-6P 703405-32-7P
     703405-33-8P 703405-34-9P 703405-35-0P
     703405-36-1P 703405-39-4P 703405-40-7P
     703405-41-8P 703405-42-9P 703405-43-0P
     703405-44-1P 703405-45-2P 703405-46-3P
     703405-47-4P 703405-48-5P 703405-49-6P
     703405-50-9P 703405-51-0P 703405-52-1P
     703405-53-2P 703405-54-3P 703405-55-4P
     703405-56-5P 703405-57-6P 703405-58-7P
     703405-59-8P 703405-60-1P
     RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT
     (Reactant or reagent)
        (preparation of N, N-bridged, nitrogen-substituted carbacyclic
        indolocarbazoles for use in pharmaceutical compns. as protein kinase
        inhibitors)
     233253-35-5 CAPLUS
RN
     9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-k1]pyrrolo[3,4-
CN
     i][1,6]benzodiazocine-1,3,10(2H,9H)-trione, 11,12-dihydro- (9CI)
     INDEX NAME)
```

Relative stereochemistry.



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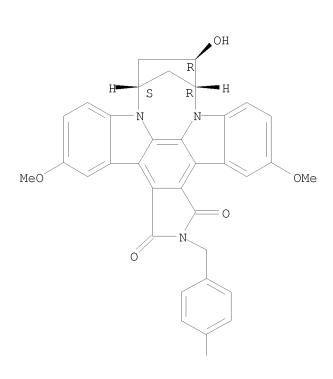
RN 253680-58-9 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3,10(2H,9H)-trione,
11,12-dihydro-2-[(4-methoxyphenyl)methyl]- (9CI) (CA INDEX NAME)

RN 703405-26-9 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carboxylic acid,
10-amino-2,3,9,10,11,12-hexahydro-1,3-dioxo-,
2-[[(1,1-dimethylethoxy)carbonyl]amino]ethyl ester, (9R,10S,12S)-rel(9CI) (CA INDEX NAME)

Relative stereochemistry.

RN 703405-29-2 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-10-hydroxy-5,16-dimethoxy-2-[(4-

methoxyphenyl)methyl]-, (9R,10R,12S)-rel- (9CI) (CA INDEX NAME)
Relative stereochemistry.



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OMe

RN 703405-30-5 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3,10(2H,9H)-trione,
11,12-dihydro-5,16-dimethoxy-2-[(4-methoxyphenyl)methyl]- (9CI) (CA INDEX NAME)

RN 703405-31-6 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3,10(2H,9H)-trione, 11,12-dihydro-5,16-dimethoxy(9CI) (CA INDEX NAME)

RN 703405-32-7 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-k1]pyrrolo[3,4-i][1,6]benzodiazocine-10-carbonitrile,
10-amino-2,3,9,10,11,12-hexahydro-5,16-dimethoxy-1,3-dioxo-,
(9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

RN 703405-33-8 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-10-carboxamide,
 10-amino-2,3,9,10,11,12-hexahydro-5,16-dimethoxy-1,3-dioxo-,
 (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

RN 703405-34-9 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-10,11-dihydroxy-2-[(4-methoxyphenyl)methyl]-,
(9R,10S,11R,12S)-rel- (9CI) (CA INDEX NAME)

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RN 703405-35-0 CAPLUS
CN 6,10-Methano-16H-[1,3,2]dioxathiolo[4,5-c]diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-16,18(17H)-dione,
 6,6a,9a,10-tetrahydro-, 8,8-dioxide, (6R,6aS,9aR,10S)-rel- (9CI) (CA INDEX NAME)

RN 703405-36-1 CAPLUS

CN 6,10-Methano-16H-diindolo[1,2,3-fg:3',2',1'-kl]oxazolo[4,5-c]pyrrolo[3,4-i][1,6]benzodiazocine-8,16,18(9H,17H)-trione, 6,6a,9a,10-tetrahydro-17-[(4-methoxyphenyl)methyl]-, (6R,6aR,9aR,10S)-rel-(9CI) (CA INDEX NAME)

Relative stereochemistry.

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0

RN 703405-39-4 CAPLUS

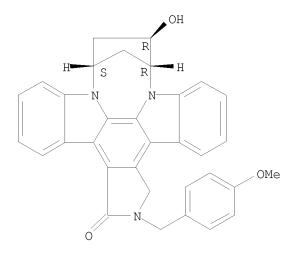
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocin-1-one, 2,3,9,10,11,12-hexahydro-11-hydroxy-2-[(4-methoxyphenyl)methyl]-, (9R,11S,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

RN 703405-40-7 CAPLUS

CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocin-1-one, 2,3,9,10,11,12-hexahydro-10-hydroxy-2-[(4-methoxyphenyl)methyl]-, (9R,10R,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.



RN 703405-41-8 CAPLUS

CN Benzeneacetic acid, 3,4-dimethoxy-, (9R,10R,12S)-2,3,9,10,11,12-hexahydro-2-[(4-methoxyphenyl)methyl]-3-oxo-9,12-methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocin-10-yl ester, rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

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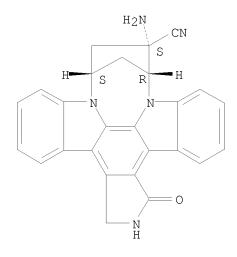
RN 703405-42-9 CAPLUS
CN Benzeneacetic acid, 3,4-dimethoxy-,
(9R,10R,12S)-2,3,9,10,11,12-hexahydro-2-[(4-methoxyphenyl)methyl]-1-oxo9,12-methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocin-10-yl ester, rel- (9CI) (CA INDEX NAME)

RN 703405-43-0 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,11(12H)-dione,
2,3,9,10-tetrahydro-2-[(4-methoxyphenyl)methyl]- (9CI) (CA INDEX NAME)

RN 703405-44-1 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-k1]pyrrolo[3,4i][1,6]benzodiazocine-1,11(12H)-dione, 2,3,9,10-tetrahydro- (9CI) (CA INDEX NAME)

RN 703405-45-2 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carbonitrile,
 10-amino-2,3,9,10,11,12-hexahydro-3-oxo-, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.



RN 703405-46-3 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,10(9H)-dione,
2,3,11,12-tetrahydro-2-[(4-methoxyphenyl)methyl]- (9CI) (CA INDEX NAME)

RN 703405-47-4 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,10(9H)-dione, 2,3,11,12-tetrahydro- (9CI) (CA INDEX NAME)

RN 703405-49-6 CAPLUS

CN 2-Propanesulfinamide, 2-methyl-N-[(9S,12R)-2,3,11,12-tetrahydro-2-[(4-methoxyphenyl)methyl]-1-oxo-9,12-methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocin-10(9H)-ylidene]-, [N(E),S(S)]- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

Double bond geometry as shown.

RN 703405-50-9 CAPLUS

CN 2-Propanesulfinamide, 2-methyl-N-[(9R,12S)-2,3,11,12-tetrahydro-2-[(4-methoxyphenyl)methyl]-1-oxo-9,12-methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocin-10(9H)-ylidene]-, [N(E),S(S)]- (9CI) (CA INDEX NAME)

Double bond geometry as shown.

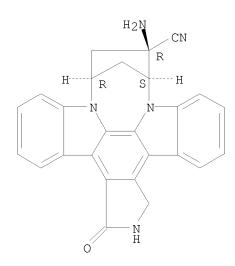
RN 703405-51-0 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,10(9H)-dione,
 2,3,11,12-tetrahydro-2-[(4-methoxyphenyl)methyl]-, (9S,12R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 703405-52-1 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,10(9H)-dione, 2,3,11,12-tetrahydro-, (9S,12R)(9CI) (CA INDEX NAME)

RN 703405-53-2 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carbonitrile,
10-amino-2,3,9,10,11,12-hexahydro-1-oxo-, (9S,10R,12R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



RN 703405-54-3 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-10-carboxamide,
10-amino-2,3,9,10,11,12-hexahydro-1-oxo-, (9S,10R,12R)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 703405-56-5 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,10(9H)-dione,
 2,3,11,12-tetrahydro-2-[(4-methoxyphenyl)methyl]-, (9R,12S)- (9CI) (CA INDEX NAME)

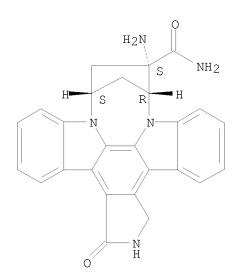
RN 703405-57-6 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,10(9H)-dione, 2,3,11,12-tetrahydro-, (9R,12S)(9CI) (CA INDEX NAME)

Absolute stereochemistry.

RN 703405-58-7 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carbonitrile,
10-amino-2,3,9,10,11,12-hexahydro-1-oxo-, (9R,10S,12S)- (9CI) (CA INDEX NAME)

RN 703405-59-8 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carboxamide,
 10-amino-2,3,9,10,11,12-hexahydro-1-oxo-, (9R,10S,12S)- (9CI) (CA INDEX NAME)

Absolute stereochemistry.



REFERENCE COUNT: 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

L6 ANSWER 7 OF 8 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 2000:31347 CAPLUS

DOCUMENT NUMBER: 132:78734

TITLE: Preparation of indolocarbazole derivatives useful for

the treatment of neurodegenerative diseases

characterized by tau hyperphosphorylation and cancer INVENTOR(S): Roder, Hanno; Lowinger, Timothy B.; Brittelli, David

R.; Vanzandt, Michael C.

PATENT ASSIGNEE(S): Bayer Corporation, USA

SOURCE: U.S., 23 pp. CODEN: USXXAM

DOCUMENT TYPE: Patent LANGUAGE: English

FAMILY ACC. NUM. COUNT: 1

PATENT INFORMATION:

PA:	PATENT NO.					KIND		DATE		APE	PLI	LICATION NO.				DATE			
US	US 6013646				A		20000111			US 1998-109131						19980702			
CA	CA 2336419				A1 20000113			CA 1999-2336419						19990623					
WO	7O 2000001699				A1 2000011		0113	WO 1999-EP4369							19990623				
	W:	ΑE,	AL,	AM,	AT,	AU,	AZ,	BA,	BB,	ВС	3,	BR,	BY,	CA,	CH,	CN,	CU,	CZ,	
		DE,	DK,	EE,	ES,	FI,	GB,	GD,	GE,	GF	ł,	GM,	HR,	HU,	ID,	IL,	IN,	IS,	
		JP,	KE,	KG,	KP,	KR,	KZ,	LC,	LK,	LF	۲,	LS,	LT,	LU,	LV,	MD,	MG,	MK,	
							PL,												
		TM,	TR,	TT,	UA,	UG,	US,	UZ,	VN,	JY	J,	ZA,	ZW						
	RW:	GH,	GM,	KE,	LS,	MW,	SD,	SL,	SZ,	UC	3,	ZW,	AT,	BE,	CH,	CY,	DE,	DK,	
		ES,	FΙ,	FR,	GB,	GR,	IE,	IT,	LU,	MC	Ì,	NL,	PT,	SE,	BF,	ВJ,	CF,	CG,	
		CI,	CM,	GA,	GN,	GW,	ML,	MR,	NE,	SI	1,	TD,	TG	•	•	·	·	,	
AU	9947766				A 20000124				·	AU 1999-47766					19990623				
AU	U 754399				B2 20021114														
EP	1091	962			A1		2001	0418		ΕP	19	99-9	9311	58		1	9990	623	
	R:	DE,	ES,	FR,	GB,	ΙT													
JP	2002	5194.	25 [.]	,	T		2002	0702		JΡ	20	00-	5581	02		1	9990	623	
US	US 6541468				В1		2003	0401	US 1999-382539						19990825				
PRIORITY APPLN. INFO.:										US	19	98-	1091	31		A 1	9980	702	
										WO	19	99-I	EP43	69	,	W 1	9990	623	
OTHER SO	THER SOURCE(S):				MARPAT		132:78734												

GI

AB Indolocarbazoles I [R1 = H, OH, carboxy, carboxamido, alkyloxyalkyl; R2, R3, R4 = H, OH; R5, R6 = H, OH, amino, acylamino, acyloxy, alkyloxy, carboxy, carboxamido, halogen; R7, R8 = H, OH, halogen; R7R8 = oxo; Z = O, H2], which are analogs of K 252a, a naturally occurring alkaloid, were prepared for potential use in the treatment of neurodegenerative diseases characterized by tau hyperphosphorylation, such as Alzheimer's disease (AD), frontal lobe degeneration (FLD), argyrophilic grains disease, subacute sclerotizing panencephalitis (SSPE), and cancer. Thus, indolocarbazole II was prepared in a 5 step synthetic sequence starting from (1R,3S)-4-cyclopentene-1,3-diol monoacetate and 12,13-dihydro-6-[(4-methoxyphenyl)methyl]-5H-indolo[2,3-a]pyrrolo[3,4-c]carbazole-5,7(6H)-dione. The prepared compds. were assayed for cAMP-dependent kinase and cdc2 kinase inhibiting activity.

IT 233253-35-5P 233253-37-7P 253680-44-3P 253680-48-7P

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological study); PREP (Preparation); USES (Uses)

(preparation of indolocarbazole derivs. useful for the treatment of neurodegenerative diseases characterized by tau hyperphosphorylation and cancer)

RN 233253-35-5 CAPLUS

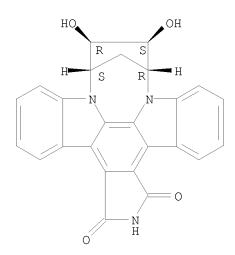
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3,10(2H,9H)-trione, 11,12-dihydro- (9CI) (CA INDEX NAME)

RN 233253-37-7 CAPLUS

CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-10-carboxylic acid, 2,3,9,10,11,12-hexahydro-10-hydroxy-1,3-dioxo-, methyl ester, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

RN 253680-44-3 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-10,11-dihydroxy-, (9R,10S,11R,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.



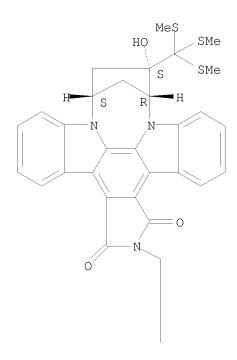
PAGE 1-A

PAGE 2-A | OMe

RN 253680-58-9 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3,10(2H,9H)-trione,
11,12-dihydro-2-[(4-methoxyphenyl)methyl]- (9CI) (CA INDEX NAME)

RN 253680-60-3 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3(2H)-dione,
9,10,11,12-tetrahydro-10-hydroxy-2-[(4-methoxyphenyl)methyl]-10[tris(methylthio)methyl]-, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)



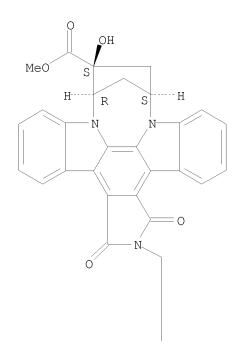


PAGE 2-A

RN 253680-62-5 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-10-carboxylic acid,
2,3,9,10,11,12-hexahydro-10-hydroxy-2-[(4-methoxyphenyl)methyl]-1,3-dioxo-, methyl ester, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

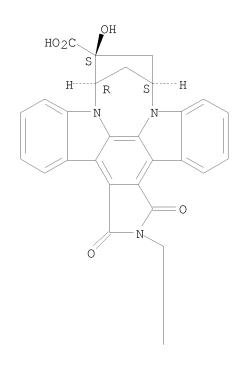
PAGE 1-A



PAGE 2-A

Relative stereochemistry.

PAGE 1-A

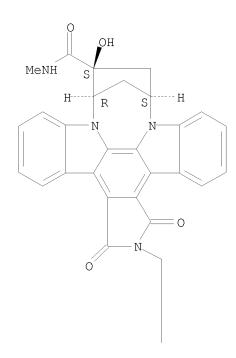


PAGE 2-A

RN 253680-66-9 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-10-carboxamide,
2,3,9,10,11,12-hexahydro-10-hydroxy-2-[(4-methoxyphenyl)methyl]-N-methyl-1,3-dioxo-, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

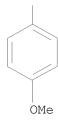
Relative stereochemistry.

PAGE 1-A



10/532,263

PAGE 2-A



REFERENCE COUNT:

17 THERE ARE 17 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT

10/532,263

AUTHOR(S):

L6 ANSWER 8 OF 8 CAPLUS COPYRIGHT 2008 ACS on STN

ACCESSION NUMBER: 1999:320447 CAPLUS

DOCUMENT NUMBER: 131:116392

TITLE: Synthesis of novel carbocyclic analogues of

indolocarbazole natural products Riley, Dean A.; Simpkins, Nigel S.

CORPORATE SOURCE: School of Chemistry, University of Nottingham,

University Park, Nottingham, NG7 2RD, UK

SOURCE: Tetrahedron Letters (1999), 40(20), 3929-3932

CODEN: TELEAY; ISSN: 0040-4039

PUBLISHER: Elsevier Science Ltd.

DOCUMENT TYPE: Journal LANGUAGE: English

OTHER SOURCE(S): CASREACT 131:116392

GΙ

AB The synthesis of some cyclopentane-bridged indolocarbazoles, such as I (R = benzyl, H) representing carbocyclic analogs of the natural product K-252a, was achieved by a concise, convergent route, and the ring expansion of one compound to a staurosporine-type derivative was also demonstrated. The products are potent inhibitors of protein kinase C (PKC) (no data).

IT 233253-26-4P 233253-27-5P 233253-28-6P 233253-30-0P 233253-31-1P 233253-34-4P

Ι

233253-35-5P 233253-36-6P

RL: RCT (Reactant); SPN (Synthetic preparation); PREP (Preparation); RACT (Reactant or reagent)

(synthesis of novel carbocyclic analogs of staurosporine and K 252a indolocarbazole natural products)

RN 233253-26-4 CAPLUS

CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-

i][1,6]benzodiazocine-1,3(2H)-dione,

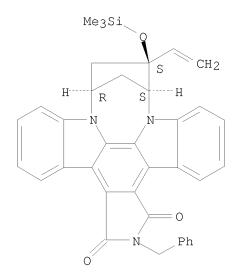
9,10,11,12-tetrahydro-10-hydroxy-2-(phenylmethyl)-, (9R,10R,12S)-rel-(9CI) (CA INDEX NAME)

RN 233253-27-5 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3,10(2H,9H)-trione,
11,12-dihydro-2-(phenylmethyl)- (9CI) (CA INDEX NAME)

RN 233253-28-6 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carbonitrile,
 2,3,9,10,11,12-hexahydro-1,3-dioxo-2-(phenylmethyl)-10[(trimethylsilyl)oxy]-, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

RN 233253-30-0 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-1,3(2H)-dione,
10-ethenyl-9,10,11,12-tetrahydro-2-(phenylmethyl)-10-[(trimethylsilyl)oxy], (9R,10R,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

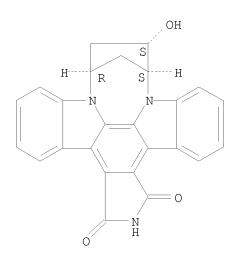


RN 233253-31-1 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-10-carboxaldehyde,
2,3,9,10,11,12-hexahydro-1,3-dioxo-2-(phenylmethyl)-10[(trimethylsilyl)oxy]-, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

RN 233253-34-4 CAPLUS

ON 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3(2H)-dione, 9,10,11,12-tetrahydro-10-hydroxy-, (9R,10R,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

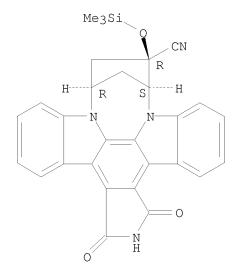


RN 233253-35-5 CAPLUS

CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-1,3,10(2H,9H)-trione, 11,12-dihydro- (9CI) (CA INDEX NAME)

RN 233253-36-6 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4i][1,6]benzodiazocine-10-carbonitrile,
2,3,9,10,11,12-hexahydro-1,3-dioxo-10-[(trimethylsilyl)oxy]-,
(9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.

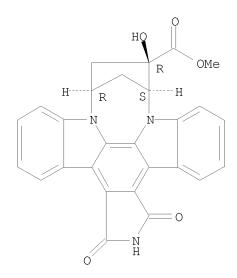


IT 233253-29-7P 233253-37-7P
 RL: SPN (Synthetic preparation); PREP (Preparation)
 (synthesis of novel carbocyclic analogs of staurosporine and K 252a
 indolocarbazole natural products)
RN 233253-29-7 CAPLUS
CN 9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4 i][1,6]benzodiazocine-10-carboxylic acid,
 2,3,9,10,11,12-hexahydro-10-hydroxy-1,3-dioxo-2-(phenylmethyl)-, methyl
 ester, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

RN 233253-37-7 CAPLUS

9,12-Methano-1H-diindolo[1,2,3-fg:3',2',1'-kl]pyrrolo[3,4-i][1,6]benzodiazocine-10-carboxylic acid, 2,3,9,10,11,12-hexahydro-10-hydroxy-1,3-dioxo-, methyl ester, (9R,10S,12S)-rel- (9CI) (CA INDEX NAME)

Relative stereochemistry.



REFERENCE COUNT:

13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD. ALL CITATIONS AVAILABLE IN THE RE FORMAT